EXHIBIT E

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BioMarin Pharmaceutical, Inc. **Biotechnology Specialist** Intellectual Property 11 Pimentel Court Rita Charles

Novato, Calif 94949

Suite 210 · Novato, CA 94949 371 Bel Marin Keys Boulevard

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Brigitte Groulx c/o_Biochem Pharma Cancer Biology Dept. 275 Armand Frappier Laval, QC H7V 4A7 Tel: 450-978-7903 Lift Shire Brochen May 2001

Re: U.S. Patent Application No. 08/722,659

Title: Use of Heparinase to Decrease Inflammatory

Response

Inventors: Bennett *et al.*Filed: September 27, 1996
Our reference No.: 104385.140

Dear Brigitte,

In furtherance of our prosecution of the above named patent, please sign the enclosed Declaration in <u>blue ink</u> and return it to me as soon as possible.

A return envelope is provided for your convenience.

If you have any questions regarding your Declaration or the present proceedings in the U.S. Patent and Trademark Office, please feel free to contact me directly at 415-884-6709 or lbigornia@bmrn.com.

Sincerely.

Luisa Bigornia Ph.D., Esq.

Attorney Docket No. 104385.140

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:)
BENNETT et al.) Group Art Unit: 1644
)
Serial No.: 08/722,659) Examiner: A. DeCloux
)
Filed: September 27, 1996)
- ·)
For: USE OF HEPARINASE TO DECR	EASE INFLAMMATORY RESPONSE

Commissioner for Patents Washington, D.C. 20231

DECLARATION UNDER 37 C.F.R. § 1.132 OF BRIGETTE GROUIX

- I, Brigette Grouix, do hereby declare that:
- I am a joint inventor of the subject matter claimed in the above-identified
 application, along with Elizabeth Cauchon, Dominique Fink, Ariane Hsia, Joseph Zimmermann,
 Clark Bennett, and Pamela Danagher.
- 2. I am informed and believe that U.S. Patent No. 5,997,863 ("the'863 patent"), a copy of which is attached as Exhibit A, has been cited against the claims of the above-identified application. In particular, I am informed and believe that Example 8 found at col. 17, line 63-col. 18, line 29 has been cited against the above-captioned application in a prior art rejection.
- 3. While I am a co-inventor of the above-captioned application, I did not make an inventive contribution to the subject matter recited in paragraph 2. In particular, I was not involved in the animal model work described in Example 8 involving the local administration of heparinase to enhance revascularization, described generally at col. 17, line 63-col. 18, line 29.
 - 4. The undersigned declares further that all statements made herein of my own

knowledge are true and that all statements made on information and belief are believed to be true and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of this application or any patent issuing therefrom.

Signed at		
		Brigette Grouix
this day of	2003	

Luisa Bigornia

DIOMAN CENTRALING

371 Bel Marin Keys Boulevard Suite 210 • Novato, CA 94949 Ariane Hsia c/o RIV/LTD.

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Ariane Hsia c/o RIV LTD. 22 Wesllesley St. East Toronto, Ontario M4Y 1Y3

> Re: U.S. Patent Application No. 08/722,659 Title: Use of Heparinase to Decrease Inflammatory

Response

Inventors: Bennett *et al.*Filed: September 27, 1996
Our reference No.: 104385.140

Dear Ariane,

In furtherance of our prosecution of the above named patent, please sign the enclosed Declaration in <u>blue ink</u> and return it to me as soon as possible.

A return envelope is provided for your convenience.

If you have any questions regarding your Declaration or the present proceedings in the U.S. Patent and Trademark Office, please feel free to contact me directly at 415-884-6709 or lbigornia@bmrn.com.

Sincerely,

Luisa Bigornia Ph.D., Esq.

LSB/rvc

Attorney Docket No. 104385.140

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of: BENNETT et al.) Group Art Unit: 1644
Serial No.: 08/722,659) Examiner: A. DeCloux
Filed: September 27, 1996)
For: USE OF HEPARINASE TO DEC) CREASE INFLAMMATORY RESPONSE

Commissioner for Patents Washington, D.C. 20231

DECLARATION UNDER 37 C.F.R. § 1.132 OF ARIANE HSIA

- I, Ariane Hsia, do hereby declare that:
- 1. I am a joint inventor of the subject matter claimed in the above-identified application, along with Elizabeth Cauchon, Dominique Fink, Brigette Grouix, Joseph Zimmermann, D. Clark Bennett, and Pamela Danagher.
- 2. I am informed and believe that U.S. Patent No. 5,997,863 ("the'863 patent"), a copy of which is attached as Exhibit A, has been cited against the claims of the above-identified application. In particular, I am informed and believe that Example 8 found at col. 17, line 63-col. 18, line 29 has been cited against the above-captioned application in a prior art rejection.
- 3. While I am a co-inventor of the above-captioned application, I did not make an inventive contribution to the subject matter recited in paragraph 2. In particular, I was not involved in the animal model work described in Example 8 involving the local administration of heparinase to enhance revascularization, described generally at col. 17, line 63-col. 18, line 29.
 - 4. The undersigned declares further that all statements made herein of my own

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knowledge are true and that all statements made on information and belief are believed to be true and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of this application or any patent issuing therefrom.

Signed at			
			Ariane Hsia
this	day of	2003	

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371 Bel Marin Keys Boulevard Sulte 210 • Novato, CA 94949



BioMarin Pharmaceutical, Inc. **Biotechnology Specialist** Intellectual Property Novato, Calif 94949 11 Pimentel Court Rita Charles

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Ms. Dominique Fink 4632 rue Chambord Montreal, Quebec H2J 3M7

Re: U.S. Patent Application No. 08/722,659

Title: Use of Heparinase to Decrease Inflammatory

Response

Inventors: Bennett *et al.*Filed: September 27, 1996
Our reference No.: 104385.140

Dear Ms. Fink,

In furtherance of our prosecution of the above named patent, please sign the enclosed Declaration in <u>blue ink</u> and return it to me as soon as possible.

A return envelope is provided for your convenience.

If you have any questions regarding your Declaration or the present proceedings in the U.S. Patent and Trademark Office, please feel free to contact me directly at 415-884-6709 or lbigornia@bmrn.com.

Sincerely,

Luisa Bigornia Ph.D., Esq.

LSB/rvc

Attorney Docket No. 104385.140

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

PA CHIMANAIN

In re A	pplication of:)	-
	BENNETT et al.)	Group Art Unit: 1644
)	•
Serial 1	No.: 08/722,659)	Examiner: A. DeCloux
)	
Filed:	September 27, 1996)	
)	
For:	USE OF HEPARINASE TO DECRE	ASE IN	VFLAMMATORY RESPONSE

Commissioner for Patents Washington, D.C. 20231

DECLARATION UNDER 37 C.F.R. § 1.132 OF DOMINIQUE FINK

- I, Dominique Fink, do hereby declare that:
- I am a joint inventor of the subject matter claimed in the above-identified
 application, along with Elizabeth Cauchon, Brigette Grouix, Ariane Hsia, Joseph Zimmermann,
 Clark Bennett, and Pamela Danagher.
- 2. I am informed and believe that U.S. Patent No. 5,997,863 ("the'863 patent"), a copy of which is attached as Exhibit A, has been cited against the claims of the above-identified application. In particular, I am informed and believe that Example 8 found at col. 17, line 63-col. 18, line 29 has been cited against the above-captioned application in a prior art rejection.
- 3. While I am a co-inventor of the above-captioned application, I did not make an inventive contribution to the subject matter recited in paragraph 2. In particular, I was not involved in the animal model work described in Example 8 involving the local administration of heparinase to enhance revascularization, described generally at col. 17, line 63-col. 18, line 29.
 - 4. The undersigned declares further that all statements made herein of my own

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knowledge are true and that all statements made on information and belief are believed to be true and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of this application or any patent issuing therefrom.

Signed at				
			Dominique Fink	
this	day of	2003.		



Ms. Dominique Fink 4632 rue Chambord Montreal, Quebec H2J 3M7

Re: U.S. Patent Application No. 08/722,659

Title: Use of Heparinase to Decrease Inflammatory

Response

Inventors: Bennett *et al.*Filed: September 27, 1996
Our reference No.: 104385.140

Dear Ms. Fink,

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Sincerely,

Luisa Bigornia Ph.D., Esq.

LSB/rvc